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SUB	STIT	JTE FORM PTO-1390 ·	U.S. DEPARTMENT OF COM PATENT AND TRADEMARK (ATTORNEY'S DOCKET NUMBER 08291-435001
		TRANSMITTAL LETTER TO DESIGNATED/ELECTED OF CONCERNING A FILING UN	FICE (DO/EO/US)		U.S. APPLICATION NO. (IF KNOWN)
					09/509308
PC	T/GE	TIONAL APPLICATION NO. 398/02863	INTERNATIONAL FILING DATE September 22, 1998		PRIORITY DATE CLAIMED September 25, 1997
DE.	ACT	INVENTION IVANTS FOR DUST MITE ALI	LERGENS		
Jan	ette	NT(S) FOR DO/EO/US SUH, Malcolm Tom McKECH	NIE, Gay CORNELIUS and	Ian Andrew	THOMPSON
App oth	olicar er int	nt herewith submits to the Unit formation:	ed States Designated/Elect	ed Office (E	OO/EO/US) the following items and
1.	\boxtimes	This is a FIRST submission o	f items concerning a filing u	nder 35 U.S	S.C. 371.
2.		This is a SECOND or SUBQU	JENT submission of items o	oncerning a	a filing under 35 U.S.C. 371.
3.	\boxtimes	This is an express request to than delay examination until the Articles 22 and 39(1).	begin national examination ne expiration of the applicat	procedures le time limi	t set in 35 U.S.C. 371(b) at any time rather t set in 35 U.S.C. 371(b) and PCT
4.	\boxtimes	A proper Demand for Internat claimed priority date.	ional Preliminary Examinati	on was mad	de by the 19th month from the earliest
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6.		A translation of amendments	to the claims under PCT Ar	ticle 19 (35	U.S.C. 371(c)(2)).
7.	\boxtimes	a. are transmitted herevb. have been transmitte	vith (required only if not tran d by the International Burea however, the time limit for	smitted by t u.	T Article 19 (35 U.S.C. 371(c)(3)) the International Bureau). h amendments has NOT expired.
8.		A translation of amendments	to the claims under PCT Ar	ticle 19 (35	U.S.C. 371(c)(3)).
9.		An oath or declaration of the			
10	. 🔲	A translation of the annexes t (35 U.S.C. 371(c)(5)).	o the International Prelimina	ary Examina	ation Report under PCT Article 36
Ite	ms 1	1. to 16. below concern other	documents or information ir	cluded:	
i		An Information Disclosure Sta			
12	. 🗆	An assignment document for 3.31 is included.	recording. A separate cover	er sheet in o	compliance with 37 CFR 3.28 and
13		A FIRST preliminary amenda A SECOND or SUBSEQUEN		"EXPRESS M	IAIL Mailing Label Number EE647186485US
14	. 🗆	A substitute specification.		Date of Depos	March 23, 2000
15		A change of power of attorne	y and/or address letter.	deposited with	y under 37 CFR 1.10 that this correspondence is being the United States Postal Service as "Express Mail
16	. ⊠	Other items or information: Copy of PCT Written Opi Copy of International Predated 17 December 1999	liminary Examination Repo	indicated abou	Addressee" with sufficient postage on the date of and is addressed to the Assistant Commissioner for ington, D.C. 20231. Clenter Figures
					Valentin Figueroa

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U.S. APPLICATION S. AFS 1993 08	INTERNATIONAL APPLIC PCT/GB98/02863	CATION NO.	ATTORNEY'S DOCKI 08291-435001		
17. The following fees are submitted	:		CALCULATIONS	PTO USE ONLY	
Basic National Fee (37 CFR 1.492(a)(1)-	(5)):				
Search report has been prepared by the	EPO or JPO	\$840	\$0.00		
International preliminary examination fee	paid to USPTO (37 0	CFR 1.482) \$670	\$0.00		
No international preliminary examination 1.482) but international search fee paid t			\$0.00		
Neither international preliminary examina international search fee (37 CFR 1.445(a			\$970.00		
International preliminary examination fee and all claims satisfied provisions of PCT			\$0.00		
ENTER	R APPROPRIATE BAS	SIC FEE AMOUNT	\$970.00		
Surcharge of \$130 for furnishing the oath	or declaration later th	han [] 20 [] 30			
mos. from the earliest claimed priority da			\$0.00		
Claims Number Filed	Number Extra	Rate	Ψ0.00		
Total Claims 16 - 20	Number Extra	x \$18	\$0.00		
Independent Claims 5- 3	2	x \$78	\$156.00		
Multiple Dependent Claims(s) (if applicat		+ \$260	\$0.00		
TOTAL OF ABOVE CALCULATIONS	JIC)	Ψ200	\$1,126.00		
Reduction by ½ for filing by small entity, if applicable. Verified Small Entity					
statement must also be filed. (Note 37 C			\$0.00		
SUBTOTAL	······································		\$1,126.00		
Processing fee of \$130 for furnishing the	e English Translation I	later than		***	
20 ☐ 30 mos. from the earliest claim			\$0.00		
TOTAL NATIONAL FEE	, , , , , , , , , , , , , , , , , , ,	3.77	\$1,126.00		
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must be accompanied by an appropriate	cover sheet (37 CFR	3.28, 3.31).	\$0.00		
TOTAL FEES ENCLOSED			\$1,126.00		
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 A check in the amount of \$1,126 Please charge my Deposit According to copy of this sheet is enclosed. The Commissioner is hereby autoverpayment to Deposit Account 	unt No. 06-1050 in the thorized to charge any	e amount of \$0.00 to o additional fees which	n may be required, c		
NOTE: Where an appropriate time limit 1.137(a) or (b) must be filed and				evive (37 CFR	
SEND ALL CORRESPONDENCE TO:			•		
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			24,488		
Date: 23 March 2000	REGISTR	ATION NUMBER			

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Applicant: Janette SUH et al. Serial No.: PCT/GB98/02863 Filed : 22 September 1998

Title

: DEACTIVANTS FOR DUST MITE ALLERGENS

BOX: PCT

Assistant Commissioner for Patents

Washington, D.C. 20231

PRELIMINARY AMENDMENT

Prior to examination, please amend the application as follows:

In the Specification:

Page 1, after the title: insert, as a heading

--Background of the Invention--.

Page 2, after line 20: insert, as a heading:

-- Disclosure of the Invention--.

In the Claims:

Claim 6, line 30 (counting each structural formula as one line): delete "xxi) urea,".

Claim 7, line 1, change "claims 1, 2, 4 or 5" to --claim 1--.

Claim 10, line 27 (counting each structural formula as one line): delete "xxi) urea,".

CERTIFICATE OF MAILING BY EXPRESS MAIL

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I hereby certify under 37 CFR §1.10 that this correspondence is being deposited with the United States Postal Service as Express Mail Post Office to Addressee with sufficient postage on the date indicated below and is addressed to the Assistant Commissioner for Patents, Washington, DC 20231

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the first than this I'm find and the start that Ŋ Applicant: Janette SUH et al. Attorney's Docket No.: 08291-435001 / 10184P1-US

Serial No.: PCT/GB98/02863 Filed: 22 September 1998

Page : 2

Claim 11, line 1: change "claims 8 or 9" to --claim 8--.

Claim 12, line 1: change "claims 8 to 11" to --claim 8--.

Claim 13, line 1: change "claims 8 to 12" to --claim 8--.

Claim 14, line 1: change "claims 8 to 13" to --claim 8--.

Claim 16, line 1: change "claims 8 to 15" to --claim 8--; same claim, line 4: change "fragramce" to --fragrance--.

In the Abstract:

Please insert the following pages 43-44 into the application.

REMARKS

Please apply any other charges or credits to Deposit Account No. 06-1050.

Respectfully submitted,

y 18 Rara

Date: 23 March 00

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Deactivants for Dust Mite Allergens

It has been known for a long time that house dust can trigger allergenic reactions in humans, such as asthma and rhinitis. It was reported, as early as 1928, that it was the dust mites in the dust that were the primary source of the allergenic response but it was only in the 1960's that researchers appreciated its significance.

It is believed that the faeces of two particular house dust mite species, Dermatophagoides farinae (known as Der-f) and Dermatophagoides pteronyssinus (known as Der-p) trigger the immune responses of the body, thereby giving rise to well known allergenic symptoms.

A review of this is given in Experimental and Applied Acarology, 10 (1991) p. 167-186 in an article 15 entitled "House dust-mite allergen" : A review by L. G. Arlian.

Both the Der-f and Der-p species are found throughout the world. In some areas, Der-f will be the sole Dermatophagoides species. In other areas Der-p will be the sole species. In still other areas, the two species are both present through, generally, one or the other will predominate.

One way to overcome these allergenic response has been to vacuum surfaces, such as carpets, that contain 2.5 the dust mites and their faeces thoroughly and often, but that is both time consuming (i.e. has to be regularly done if one wants to make an allergenic free environment) and is very dependant on the efficiency of vacuum cleaner 30 and filter bag used e.g. micron filter bag or 2-layer vacuum bags.

25

An alternative method of creating an allergen-free environment has been to denature the allergen, for example as described in US Patent No. 4,806,526. The only effective method however of which we are aware is to apply tannic acid to the allergen. However, tannic acid can cause staining, and this is a particularly acute problem for light coloured carpets (e.g. white and light beige carpets) and other textile surfaces as tannic acid leaves a deep brown stain.

Therefore, we have been looking for allergenic denaturants which will not stain susceptible surfaces such as carpets and still deactivate the allergen.

We have discovered a number of allergen deactivants which are effective against both the Der-f and the Der-p species. Quite surprisingly, we have discovered that some of these deactivants are specific to the type of dust mite allergen being treated. For example an effective Der-f allergen deactivants will not automatically work effectively as a Der-p allergen deactivant.

According to the invention there is provided a method for deactivating allergens derived from the Der-f and/or Der-p dust mite species, which comprises contacting the allergen with a deactivating effective amount of one or more of deactivants (herein after defined as the deactivant).

The deactivants effective against one or both of Der-f allergens and Der-p allergens are:

- i) cedarwood oil,
- 30 ii) hexadecyltrimethylammonium chloride,
 - iii) aluminium chlorohydrate,
 - iv) 1-propexy-propanol-2,
 - v) polyquaternium-10

	vi)	silica gel,
	vii)	propylene glycol alginate,
	viii)	ammonium sulphate,
	ix)	hinokitiol,
5	x)	L-ascorbic acid,
	xi)	"immobilised tannic acid", (hereinafter
		defined)
	xii)	chlorohexidine,
	xiii)	maleic anhydride,
10	xiv)	hinoki oil,
	xv)	a composite of AgCl and TiO _{2,}
	xvi)	diazolidinyl urea,
	xvii)	6-isopropyl-m-cresol,
	xviii)	a compound of formula I
		O

xix) the compound of formula II

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xx) a polymeric dialdehyde containing two or more of a recurring unit of the formula III

where n = 2 to 200,

25

xxi)

urea,

xxii) cyclodextrin,

xxiii) hydrogenated hop oil,

xxiv) polyvinylpyrrolidone,

10 xxv) N-methylpyrrolidone,

xxvi) the sodium salt of anthraquinone,

xxvii) potassium thioglycolate, and

xxviii) glutaraldehyde

Deactivants (i) through (xx) are effective against both
Der-f and Der-p allergens. Deactivants (xxi) through
(xxvi) are effective against Der-f allergens only.

Deactivants (xxvii) and (xxviii) are effective against
Der-p allergens only.

A compound of formula I is commercially available as 20 Aerosol OT.

The compound of formula II is commercially available as parsley camphor.

Hinoki oil is a mixture of thujan-3-one, 2-pinene, 3,5,7,3',4'-pentahydroflavanone and 1,3,3-trimethyl-2-norcamphanone.

Hydrogenated Hop Oil is the potassium salt of tetrahydroiso humulinic acid (also known as reduced isomerised hop extract).

Propylene glycol alginate is

5

Chlorohexadine is 1,1'-hexamethylenebis[5-(4-chlorophenyl)]-biguanide.

Hinokitol is $\beta\text{-thujaplicin},\ a\ compound\ of\ the$ formula

10

Germall II is diazolidinylurea.

Thymol is 6-isopropyl-m-cresol.

Cedarwood oil contains $\alpha-$ and $\beta-$ cedrene (ca 80%), cedrol (3-14%) and cedrenol. Other sesquiterpenes and some monoterpenes are also present.

20

30

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Polyquaternium-10 is a polymeric quaternary ammonium salt of hydroxyethyl cellulose reacted with a trimethyl ammonium substituted epoxide commercially available as Polymer JR-125.

5 Silica gel is also known as colloidal silica or silicic acid and is commercially available as Kent.

"Immobilised tannic acid" is tannic acid on polyvinyl pyrrolidone beads. Immobilised Tannic Acid was prepared as follows: 100 mg of tannic acid was dissolved in water; 50 mg of Polyclar 10 (ISP, Guildford Surrey) polyvinyl pyrrolidone beads were added and stirred for one hour; the beads were filtered off the solution and washed with a few mls of iced water until no colour was seen in the washings; they were then dried in the oven at 50°C. 15

The composite of silver chloride and TiO, is made up of 20% wt/wt AgCl on 80% TiO, 3-5 μm porous beads.

In compositions containing the deactivant, the deactivant is present in an amount of from 0.01% to 7%, preferably from 0.01% to 3%.

In methods for treating rugs and carpets to deactivate allergents, the amount of deactivant present is from about 16gm to about 170gm per 10 square meters, preferably about 32gm per 10 square meters.

25 Preferably the deactivant is selected from

xiii) maleic anhydride,

xiv)	hinoki oil,
xv)	a composite of AgCl and ${\rm TiO_2}$,
xvi)	diazolidinyl urea
xvii)	6-isopropyl-m-cresol,
xii)	chlorohexidine,

xxvi) the sodium salt of anthraquinone and xviii) a compound of formula I or II, defined above, and xix) a compound of formula II, defined above.

Further according to the invention there is provided an aerosol composition containing

- i) cedarwood oil,
- ii) hexadecyltrimethylammonium chloride,
- iii) aluminium chlorohydrate,
- iv) 1-propoxy-propanol-2,
 - v) polýquaternium-10
 - vi) silica gel,
 - vii) propylene glycol alginate,
 - viii) ammonium sulphate,
- ix) hinokitiol,
 - x) L-ascorbic acid,
 - xi) "immobilised tarnic acid", (hereinafter
 defined)
 - xii) chlorohexidine,
- 20 xiii) maleic anhydride,
 - xiv) hinoki oil,
 - xv) a composite of AgCl and TiO,
 - xvi) diazolidinyl urea,
 - xvii) 6-isopropyl-m-crescl,
- 25 xviii) a compound of formula I

$$N_{a_3} \bigoplus_{O} O O Ctyl$$

The control of the co

xix) the compound of formula II

a polymeric dialdehyde containing two or more of a recurring unit of the formula III

where n = 2 to 200, xxi) urea, xxii) cyclodextrin, hydrogenated hop oil, 10 xxiii) xxiv) polyvinylpyrrolidone, N-methylpyrrolidone, (vxx the sodium salt of anthraquinone, xxvi) xxvii) potassium thioglycolate, and 15 xxviii) glutaraldehyde

- b) a propellant, and
- c) optionally, a solvent.

Preferably the amount of deactivant present in such a composition is from 0.01% to 7%, more preferably 0.01% to 3%,

Preferably the amount of propellant present in such a composition is from 4% to 50%, more preferably from 4% to 30%,

Preferably the amount of solvent present in such a composition is 0% to 99.95%, more preferably 0% to 90%, and most preferably from 20% to 90%.

10 Preferably the deactivant in such aerosol composition is selected from

hinoki oil,
a composite of AgCl with TiO2,
diazolidinyl urea,
6-isopropyl-m-cresol,
chlorohexidine,
maleic anhydride,
the sodium salt of anthraguirone, and
a compound of formula I or II defined above.

Preferably the propellant is selected from those commercially available, for example $C_{1,4}$ alkanes, chlorofluorocarbons and compressed gases such as nitrogen, air and carbon dioxide.

Preferably the solvent is selected from $C_{\text{1-6}}$ alcohols (e.g. ethanol) or water.

In addition, the compositions of this invention may also contain one or more of the following:

a fragrance, preferably in an amount of 0% to 5%, more preferably 0% to 2%;

an antimicrobial compound e.g. alkyldimethylbenzyl ammonium saccharinate, preferably in an amount of 0.01% to 1%;

- a surfactant, e.g. Dow Corning 193 Surfactant, preferably in an amount of 0.01% to 1%;
 - a corrosion inhibitor, e.g. sodium nitrite, sodium benzoate, triethanolamine and ammonium hydroxide, preferably in an amount of 0.01% to 10%; and
- a miticide, such as benzyl benzoate, pyrethroid pemethrin, d-allethrin and optionally a synergist such as piperonyl butoxide, preferably in an amount of 0.1% to 10%.
- It has been found that deactivants of the invention 15 have as effective allergen deactivating properties as tannic acid but without the drawback of staining.

The invention will now be illustrated by the following Examples.

Examples

The test procedure in Examples 1 to 55 is as follows and is known as the ELISA protocol.

The ELISA protocol for Der-f and Der-p has been developed as follows as a measure of denaturing property for denaturants.

25 ELISA Protocol 1

 Dust is collected from Hoover™ vacuum cleaner bags and passed through a series of sieves down to 63 microns.

- 2. Clean petri dishes are labelled with the chemical to be tested (on the base). Three replicates are used for each treatment.
- 3. Filter paper is used to line each dish and 0.2g of dust is added to each dish onto the filter paper. The lid (or base, as dishes are inverted) is replaced and the dish is shaken to disperse the dust evenly over the filter paper.
- 4. 2% aqueous solutions of deactivant were used except

 for the silver chloride composite where 0.05% was used

 instead. Immobilised tannic acid was used as a 1%

 dispersion. The hydrogenerated hop end was used at the

 2% level (in the form of a 10% solution). Water
 insoluble deactivants were emulsified with a sorbitone

 oleate surfactant (i.e. Tween). Hinokitol was used at

 0.5% not 2%.
 - 5. The dust is sprayed with the corresponding treatment, 2 sprays are required for sufficient coverage(1 spray = 1.5 ml).
- 20 6. Leave uncovered at room temperature, in well aerated room, until filter paper is dry. This can take up to 4 hours.
 - 7. Empty dust in epindorfs labelled according to treatment.
- 25 8. Add 1 ml of 5% Bovine Serum Albumen Phosphate Butter Saline - Tween BSA-PBS-T to each epindorf (5 times the weight of dust) (20ml of BSA-PBS-T =1 g of BSA in 20ml of PBS-T).
 - 9. Leave overnight in a refrigerator.
- 30 10. Centrifuge for 5 minutes at 13,000 rpm.

- 11. Decant the supernatant into a new epindorf labelled according to treatment.
- 12. Centrifuge again for 5 minutes at 13,000 rpm.
- 13. Make up dilutions of 1:10 and 1:100 by adding 100 μ l of neat solution to 900 μ l of 1% BSA-PBS-T (1:10). This is repeated using 100 μ l of 1:10 dilution and add to 900 μ l of 1% BSA-PBS-T for 1:100 dilution.

ELISA Protocol 2 for Der-f and Der-p: Indoor Biotechnologies

- 10 1. Prepare samples and dilutions as in protocol
 - 2. Prepare 500 ml of 50 mM carbonate/bicarbonate buffer by dissolving 0.795g Na_2CO_3 and 1.465g $NaHCO_3$ in 500 ml of distilled water. Check the pH is at 9.6. (This solution is kept in the refrigerator in a conical flask).
- 15 3. Monoclonal antibody (kept in the freezer) has to be added to the buffer using the following method, (1 μ g per well; 11 ml is needed) applied to the ELISA plate
 - 11ml of carbonate/bicarbonate buffer is added to the dispensing tray.
- 11μl of Der-fl or Der-pl monoclonal antibody

(Stored in freezer, epindorf in use is in the refrigerator) is added to the buffer. To ensure that all the antibody is removed from the tip, wash out the pipette tip by sucking up and down I the buffer solution, gently stirring to mix thoroughly.

4. Pipette 100 μ l of the antibody solution into each well of the microtiter plate, cover with a plate sealer and leave overnight at 4°C.

- 5. Empty the plate by quickly inverting it over the sink, then dry by banging on a stack of paper towels.
- 6. Add 200 μ l of wash buffer to each well: PBS/0/05% tween (PBS-T).
- 7. Repeat stages 5 and 6 once more (making a total of 2 washes).
 - 8. Make sure all the wells are dry, then add 100 μl of 1% BSA-PBS-T. Replace the plate sealer and incubate for 1 hour at room temperature*.
- 10 9. Repeat steps 5 to 7 (2 washes).
 - 10. *During the hour incubation period, prepare the allergen standards at dilutions between 125 and 1 μ g/ml Der f 1 or Der p1:
- Add 25 μ l of allergen standard (kept in the refrigerator in polystyrene box) to 475 μ l of 1% PBS-BSA-T and mix thoroughly labelled '125'.
 - 250 μ l of 1% PBS-BSA-T is added to 7 further epindorfs which are labelled 62.5, 31.25, 15.63, 7.61, 3.9, 1.95 and 0.98.
- 20 250 μ l is taken from the 1st epindorf (labelled 125) and transferred to the next (labelled 62.5). This is mixed thoroughly.
- Using a new pipette tip, 250 μl is removed from epindorf labelled 62.5 and transferred to 31.25,
 this procedure is continued down to the 0.98 concentration (125, 62.5, 31.25, 15.63, 7.61, 3.9, 1.95, 0.98)
 - In total $475 + (250 \times 7) = 2.3 \text{ml} : 0.023 \text{g of}$ BSA added to 2.3 ml of PBS-T.

- 11. Add $100\mu l$ aliquots of the allergen sample to the plate along with the standard allergen samples for the reference curve in duplicate. The standards usually go in the first two columns on the left hand side, with the least concentrated on top. Incubate for 1 hour.
- 12. Follow stages 5 to 6, completing a total of 5 washes.
- 13. Pour 11 ml of 1% BSA-PBS-T(0.11g of BSA to 11ml of PBS-T) to the dispensing tray. Add 11 μ l of the biotinylated monoclonal antibody (refrigerator) and mix thoroughly.
 - 14. Pipette 100 μl into each well and incubate for 1 hour at room temperature.
- 15. Empty plate and wash as described in stage 12. (5 washes).
 - 16. Add 11 μ l of Streptavidin (freezer) to 11 ml of 1%BSA-PBS-T. Pipette 100 μ l into each well and incubate for 30 minutes. Reserve any remaining solution in a vial.
- 20 17. Empty plate and wash as described in stage 12 (5 washes).
 - 18. Make a solution of OPD, by putting the two tablets (in silver and gold foil) into 20 ml of distilled water (in a glass vial). Shake quite vigorously in the dark
 - 5 until the tablets have dissolved (Wrap the vial up either in tin foil or paper towel).
 - 19. Add a small amount to the remaining solution from stage 16. Wait for a colour change (positive reaction). Add 200 μl to each well and incubate for a minimum of 30 minutes in the dark.

20. Read the plate at 450nm/405nm if filter not available.

Examples 1 to 26

The deactivants, as set out in the following table,

were used against Der-f allergens according to the above procedure and the results are as given below. Tannic acid was used as a comparator. What was measured after treatment with deactivant and tannic acid was the amount of allergen remaining active after treatment. The ratio of amount of remaining active allergen after treatment with deactivant and tannic acid is also given.

Table

Example	Deactivant	Amount of Allergen remaining active after	Amount of Allergen	Ratio of remaining Number active allergen	Number
		deactivant treatment	remaining active after tannic acid	after Deactivant/Tannic	
			treatment	Acid Treatment	
		3750	1500	2.500	xxi
	Urea	1325	550	2.409	XX
2	Polymeric dialdehyde	0001	750	2.400	
3	Cedarwood oil	0001	0021	2 265	XXII
4	Cyclodextrin	3850	0001	1966	
	hexadecyltrimethylammonium chloride	4075	0081		
	Al	1675	750		13
9	Aluminali cinoloni yarate	3950	0081	2.194	Λ
7	1-propoxy-propanol-2	2027 5	933.5	2.183	. I.
∞	Silica Gel (Kent)	7335	2000	2.168	^
6	polyquatemium-10 (Polymer JR-125)	1100	055	2.000	xxiii
10	Hydrogenated Hop Oil	1100	1000	898 1	
-	Pronviene glycol alginate	3175	00/1	1.000	
	Delicities and idone	2450	1425		
71	Fully villy) pytrolidolic	2750	1700	1.618	viii
13	Ammonium sulphate				

Fyamnle	Deactivant	Amount of Allergen	Amount of	Ratio of remaining Number	Number
Tyanın hıcı		remaining active after	Allergen	active allergen	
		deactivant treatment	remaining active	after	
			after tannic acid	Deactivant/Tannic	
			treatment	Acid Treatment	
17	Hinakital (0.5%)	3065	2000	1.533	ix
71	M methyl myrrolidone	1600	1175	1.362	XXV
17	1 Assorbis Asid	2000	1500	1.333	×
17	Immobilised Tannic Acid	1550	1175	1.319	xi
\		1525	1175	1.298	xviii
× :	Aerosol O I	1412.5	1425	166.0	xii -
61	Dordon Complex	1225	1387.5	0 883	xix
707	Falsicy Campion	1312.5	1500	0.875	XIII
17	Maleic annyunuc	1530	2000	0.765	xxvi
77	Hindi oil	1025	1387.5	0.739	xiv
22	Composite of AgCl and TiO,	1025	1425	0.719	XV
36	Gammill II	950	1387.5	0.685	xvi
26	Thymol	725	1387.5	0.523	xvii
21					

Examples 27 to 47

The deactivants, as set out in the following table, were used against Der-p allergens according to the above procedure and the results are as given below. What was measured were the amount of allergens remaining after treatment with deactivant and the amount of allergens remaining after vacuuming with no deactivant treatment.

Table

Example	Deactivant	Amount of active Allergen	Amount of active	Deactivant
•		remaining after deactivant	Allergen remaining after	
		treatment	no deactivant treatment	
			but only vaccuming	
	Glutaraldehyde	816	3375	xxviii
2	Polymeric dialdehyde	2792	3375	xx
6	Cedarwood oil	3375	0009	
4	hexadecyltrimethylammonium chloride	2863	4992	ij
5	Aluminium chlorohydrate	978	4992	111
9	1-propoxy-propanol-2	1233	4992	iv
7	Silica Gel (Kent)	1540	4992	vi
8	polyquaternium-10 (Polymer JR-125)	5463	6250	^
6	Propylene glycol alginate	3781	6250 vii	vii
10	Ammonium sulphate	2325	6250	viii
	Potassium thioglycolate	3092	3375 xxvii	xxvii

Example	Deactivant	Amount of active Allergen remaining after deactivant treatment	Amount of Allergen remaining after no deactivant	Deactivant
			treatment	
1.0	Hinakital (1) 5%)	2058	3375	ix
7 .	Linkhitich (5,2,2)	1438	5642	×
1.3	L-Ascolute Acid	1125	5642	xi
4	IMMODILISEU TAILING ANN	4494	5642 xviii	xviii
CI	Aerosol U1	2281	4450	xii
16	Chlorohexidine	1030	7450	viv
17	Parsley Camphor	1007	0044	VIV
18	Maleic anhydride	/83	4430 XIII	XIII ·
10	Hinoki oil	1644	3400	XIV
20	Composite of AuCl and TiO.	1632	3400	۸x
24		1500	3400	xvii
17	1111711101			

Examples 48-55

Further samples were tested as above and compared against tannic acid. The ratio of actives remaining after deactivant treatment and actives remaining after tannic acid treatment are given below:

Example	Deactivant	atio of actives remaining after deactivant treatment over those remaining after tannic acid treatment	Number
48	Germall II	1.5	vi
49	N-methyl pyrrolidone	4.0	xv
50	Hinoki Oil	4.0	iv
51	Silver chloride/TiO2	3.5	v
52	Thymol	4.0	Vii
53	Chlorohexidine	3.0	ii
54	Maleic anhydride	1.0	iii
55	Glutaraldehyde	1.5	xviii

Examples 56-59

The following formulations can be made up as carrier compositions for use in an aerosol for deactivating Der-f and Der-p allergens.

Raw Ingredient Description By Weight	Item Classification	<u>&</u>
Anhydrous Ethanol (SD Alcohol 40)	Solvent	79.646
Alkyl dimethyl benzyl ammonium saccharinate	Cationic Surfactant	0.106
Corrosion Inhibitor (I)		0.192
Corrosion Inhibitor (II)		0.192
Corrosion Inhibitor (III)		0.096
Deionized Water	Water/Solvent	15.768
Carbon Dioxide	Propellant	4.000
TOTAL		100.000

Raw Ingredient	Them 01	10
	Item Classification	%
Description by Weight		
Anhydrous Ethanol (SD	Solvent	* 57.000
Alcohol 40)]
Fragrance#17	Fragrance	0.0500
		0.0300
Dow Corning 193	Surfactant	0.025
Surfactant		}
Corrosion Inhibitor (I)		0.100
Corrosion Inhibitor (II)		0.100
Deionized Water	Water/solvent	* 14.725
NP-40/Butane 40	Hydrocarbon	28.000
	propellant	
TOTAL		100.000

^{* =} May replace with 95% Ethanol (SD Alcohol 40) at 61.755% by weight and 9.970% by weight Deionized water

<u>Raw Ingredient</u> <u>Description by Weight</u>	Item Classification	<u>%</u>
Anhydrous Ethanol (SD Alcohol 40)	Solvent	79.646
Benzyl Benzoate - an acaricide	Active/ester	4.600
Alkyl dimethyl benzyl ammonium saccharinate	Cationic Surfactant	0.106
Corrosion Inhibitor(I)		0.192
Corrosion Inhibitor (II)		0.192
Corrosion Inhibitor (III)		0.096
Deionized Water	Water/solvent	11.168
Carbon Dioxide	Propellant	4.000
TOTAL		100.000

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Raw Ingredient	Item Classification	8
Description by weight	TECM CIASSILICACION	30
Description by Weight		1
Anhydrous Ethanol (SD	Solvent	*57.000
Alcohol 40)		
Benzyl Benzoate	Active/ester	4.600
Demay 1 Demadace	ACCIVE/ estel	4.600
Fragrance#17	Fragrance	0.0500
Dow Corning 193	Surfactant	0.025
Surfactant	_	
Corrosion Inhibitor (I)		0.700
COLLOSION INNIBITOR (1)		0.100
Corrosion Inhibitor (II)		0.100
Deionized Water	Water/solvent	*10.125
NP-40/Butane 40	Hydrocarbon	28.000
40/ Ducaire 40		28.000
	propellant	
TOTAL		100.000

^{* =} May replace 95% Ethanol (SD Alcohol 40) at 61.755% by weight and 5.370% by weight Deionized water.

CLAIMS

1. A method for deactivating a Der-f and/or a Der-p allergen comprising contacting the allergen with a deactivating effective amount of one or more of deactivants selected from

- i) cedarwood oil, ii) hexadecyltrimethylammonium chloride, iii) aluminium chlorohydrate, iv) 1-propoxy-propanol-2, v) polyguaternium-10 vi) silica gel, vii) propylene glycol alginate, ammonium sulphate, viii) ix) hinokitiol. x) L-ascorbic acid, xi) immobilised tannic acid, xii) chlorohexidine. xiii) maleic anhydride, xiv) hinoki oil, xv) a composite of AgCl and TiO, diazolidinyl urea, xvi) xvii) 6-isopropyl-m-cresol, xviii) a compound of formula I
 - O octyl
 Na₃ OS O octyl

the compound of formula II xix)

xx)a polymeric dialdehyde containing two or more of a recurring unit of the formula III

where n = 2 to 200, xxi) urea,

> xxii) cyclodextrin,

xxiii) hydrogenated hop oil,

xxiv) polyvinylpyrrolidone,

(vxx N-methylpyrrolidone,

xxvi) the sodium salt of anthraquinone,

xxvii) potassium thioglycolate, and

glutaraldehyde. xxviii)

- A method for deactivating a Der-f allergen comprising contacting the allergen with a deactivating effective amount of one or more deactivants selected from
 - i) cedarwood oil,
 - ii) hexadecyltrimethylammonium chloride,

iii)	aluminium chlorohydrate,
iv)	1-propoxy-propanol-2,
v)	polyquaternium-10
vi)	silica gel,
vii)	propylene glycol alginate,
viii)	ammonium sulphate,
ix)	hinokitiol,
x)	L-ascorbic acid,
xi)	immobilised tannic acid,
xii)	chlorohexidine,
xiii)	maleic anhydride,
xiv)	hinoki oil,
xv)	a composite of AgCl and TiO_2
xvi)	diazolidinyl urea,
xvii)	6-isopropyl-m-cresol,
xviii)	a compound of formula I

xix) the compound of formula II

xx) a polymeric dialdehyde containing two or more of a recurring unit of the formula III

where n = 2 to 200,

xxi)

urea,

xxii)

cyclodextrin,

xxiii)

hydrogenated hop oil,

xxiv)

polyvinylpyrrolidone,

xxv)

N-methylpyrrolidone, and

xxvi)

the sodium salt of anthraquinone.

- A method for deactivating a Der-p allergen comprising contacting the allergen with a deactivating effective amount of one or more deactivants selected from
 - i) cedarwood oil,
 - ii) hexadecyltrimethylammonium chloride,
 - iii) aluminium chlorohydrate,
 - iv) 1-propoxy-propanol-2,
 - V) polyquaternium-10
 - vi) silica gel,
 - vii) propylene glycol alginate,
 - viii) ammonium sulphate,
 - ix) hinokitiol.
 - \mathbf{x}) L-ascorbic acid,
 - xi) immobilised tannic acid.
 - xii) chlorohexidine,
 - xiii) maleic anhydride,

xiv) hinoki oil,

xv) a composite of AgCl and TiO₂.

xvi) diazolidinyl urea,

xvii) 6-isopropyl-m-cresol,

xviii) a compound of formula I

$$O$$
 octyl O octyl O octyl O

xix) the compound of formula II

a polymeric dialdehyde containing two or
more of a recurring unit of the
formula III

where n = 2 to 200,

xxvii) potassium thioglycolate, and

xxviii) glutaraldehyde.

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4. A method for deactivating allergens deriving from Der-f and/or Der-p dust mites, said allergens being associated with faecal particles excreted by said mites on the surfaces of fabric materials selected from rugs, carpet and upholstered furniture, which method comprises applying to said fabric materials a deactivant selected from

- i) cedarwood oil, ii) hexadecyltrimethylammonium chloride, iii) aluminium chlorohydrate, iv) 1-propoxy-propanol-2, v) polyquaternium-10 vi) silica gel, vii) propylene glycol alginate, viii) ammonium sulphate, ix) hinokitiol, \mathbf{x}) L-ascorbic acid, xi) immobilised tannic acid, xii) chlorohexidine, xiii) maleic anhydride, xiv) hinoki oil, xv) a composite of AgCl and TiO2 xvi) diazolidinyl urea, xvii) 6-isopropyl-m-cresol, xviii) a compound of formula I

xix) the compound of formula II

$$O \longrightarrow OCH_3$$

$$O \longrightarrow OCH_3$$

xx) a polymeric dialdehyde containing two or more of a recurring unit of the formula III

where n = 2 to 200,

xxi) urea,

xxii) cyclodextrin,

xxiii) hydrogenated hop oil,

xxiv) polyvinylpyrrolidone,

xxv) N-methylpyrrolidone,

xxvi) the sodium salt of anthraquinone,

xxvii) potassium thioglycolate, and

xxviii) glutaraldehyde

at an application rate of from 16 grams to 170 grams of deactivant per 10 square meters.

5. A method according to claim 4 in which the allergens derive from Der-f dust mites and the deactivant is selected from

i)	cedarwood oil,
ii)	hexadecyltrimethylammonium chloride,
iii)	aluminium chlorohydrate,
iv)	1-propoxy-propanol-2,
v)	polyquaternium-10
vi)	silica gel,
vii)	propylene glycol alginate,
viii)	ammonium sulphate,
ix)	hinokitiol,
x)	L-ascorbic acid,
xi)	immobilised tannic acid,
xii)	chlorohexidine,
xiii)	maleic anhydride,
xiv)	hinoki oil,
xv)	a composite of AgCl and TiO_{2}
xvi)	diazolidinyl urea,
xvii)	6-isopropyl-m-cresol,
xviii)	a compound of formula I

xix) the compound of formula II

 xx) a polymeric dialdehyde containing two or
more of a recurring unit of the
formula III

where n = 2 to 200,

- xxi) urea
- xxii) cyclodextrin,
- xxiii) hydrogenated hop oil,
- xxiv) polyvinylpyrrolidone,
- xxv) N-methylpyrrolidone, and
- xxvi) the sodium salt of anthraquinone.
- 6. A method according to claim 4 in which the allergens derive from Der-p dust mites and the deactivant is selected from
 - i) cedarwood oil,
 - ii) hexadecyltrimethylammonium chloride,
 - iii) aluminium chlorohydrate,
 - iv) 1-propoxy-propanol-2,
 - v) polyquaternium-10
 - vi) silica gel,
 - vii) propylene glycol alginate,
 - viii) ammonium sulphate,
 - ix) hinokitiol,
 - x) L-ascorbic acid,
 - xi) immobilised tannic acid,
 - xii) chlorohexidine,
 - xiii) maleic anhydride,

xiv) hinoki oil,

xv) a composite of AgCl and TiO2.

xvi) diazolidinyl urea,

xvii) 6-isopropyl-m-cresol,

xviii) a compound of formula I

$$\begin{array}{c} O \\ O \\ O \end{array} \begin{array}{c} O \\ O \end{array}$$

xix) the compound of formula II

xx) a polymeric dialdehyde containing two or more of a recurring unit of the formula III

where n = 2 to 200,

xxi) urea,

xxvii) potassium thioglycolate, and

xxviii) glutaraldehyde.

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7. A method according to any of claims 1, 2, 4 or 5 in which the deactivant is selected from

```
xiv) hinoki oil,
```

- xv) a composite of AgCl with TiO2,
- xvi) diazolidinyl urea
- xvii) 6-isopropyl-m-cresol,
- xii) chlorohexidine,
- xiii) maleic anhydride,
- xxvi) the sodium salt of anthraquinone,
- xviii) a compound of formula I, and
- xix) the compound of formula II.
- 8. An aerosol composition containing
 - a) a deactivant selected from
 - i) cedarwood oil,
 - ii) hexadecyltrimethylammonium chloride,
 - iii) aluminium chlorohydrate,
 - iv) 1-propoxy-propanol-2,
 - v) polyquaternium-10
 - vi) silica gel,
 - vii) propylene glycol alginate,
 - viii) ammonium sulphate,
 - ix) hinokitiol,
 - x) L-ascorbic acid,
 - xi) immobilised tannic acid,
 - xii) chlorohexidine,
 - xiii) maleic anhydride,
 - xiv) hinoki oil,
 - xv) a composite of AgCl and TiO,
 - xvi) diazolidinyl urea,
 - xvii) 6-isopropyl-m-cresol,

xviii) a compound of formula I

xix) the compound of formula II

xx) a polymeric dialdehyde containing two or more of a recurring unit of the formula III

where n = 2 to 200,

xxi) urea,

xxii) cyclodextrin,

xxiii) hydrogenated hop oil,

xxiv) polyvinylpyrrolidone,

xxv) N-methylpyrrolidone,

xxvi) the sodium salt of anthraquinone,

xxvii) potassium thioglycolate, and

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xxviii) glutaraldehyde;

- b) a propellant; and
- c) optionally, a solvent.
- 9. An aerosol composition according to claim 8 in which the deactivant is selected from
 - i) cedarwood oil,
 - ii) hexadecyltrimethylammonium chloride,
 - iii) aluminium chlorohydrate,
 - iv) 1-propoxy-propanol-2,
 - v) polyquaternium-10
 - vi) silica gel,
 - vii) propylene glycol alginate,
 - viii) ammonium sulphate,
 - ix) hinokitiol,
 - x) L-ascorbic acid,
 - xi) immobilised tannic acid,
 - xii) chlorohexidine,
 - xiii) maleic anhydride,
 - xiv) hinoki oil,
 - xv) a composite of AgCl and TiO₂.
 - xvi) diazolidinyl urea,
 - xvii) 6-isopropyl-m-cresol,
 - xviii) a compound of formula I

xix) the compound of formula II

a polymeric dialdehyde containing two or
more of a recurring unit of the
formula III

where n = 2 to 200,

xxi) urea,

xxii) cyclodextrin,

xxiii) hydrogenated hop oil,

xxiv) polyvinylpyrrolidone,

xxv) N-methylpyrrolidone, and

xxvi) the sodium salt of anthraquinone.

- 10. An aerosol composition according to claim 8 in which the deactivant is selected from
 - i) cedarwood oil,
 - ii) hexadecyltrimethylammonium chloride,
 - iii) aluminium chlorohydrate,
 - iv) 1-propoxy-propanol-2,
 - v) polyquaternium-10

vi)	silica gel,			
vii)	propylene glycol alginate,			
viii)	ammonium sulphate,			
ix)	hinokitiol,			
x)	L-ascorbic acid,			
xi)	immobilised tannic acid,			
xii)	chlorohexidine,			
xiii)	maleic anhydride,			
xiv)	hinoki oil,			
xv)	a composite of AgCl and TiO_2			
xvi)	diazolidinyl urea,			
xvii)	6-isopropyl-m-cresol,			
xviii)	a compound of formula I			

xix) the compound of formula II

a polymeric dialdehyde containing two or
more of a recurring unit of the
formula III

where n = 2 to 200,

xxi) urea,

xxvii) potassium thioglycolate, and

xxviii) glutaraldehyde.

11. A composition according to claims 8 or 9 in which the deactivant is selected from

xiv) hinoki oil,

xv) a composite of AgC1 with TiO₂,

xvi) diazolidinyl urea

xvii) 6-isopropyl-m-cresol,

xii) chlorohexidine.

xiii) maleic anhydride,

xxvi) the sodium salt of anthraquinone,

xviii) a compound of formula I, and

xix) the compound of formula II.

12. A composition according to any of claims 8 to 11 in which the amount of deactivant present is from 0.01% to 7%, the amount of propellant present is from 0.05% to 3%, and the amount of solvent present is from 0% to 99.95%, all percentages being by weight.

- 13. A composition according to any one of claims 8 to 12 in which the propellant is selected from C_{14} alkane and carbon dioxide.
- 14. A composition according to any one of claims 8 to 13 in which the solvent is selected from C_{1-6} alcohols or water.
- 15. A composition according to claim 14 in which the solvent is ethanol.
- 16. A composition according to any one of claims 8 to 15 in which the composition may also contain one or more of the following:
 - a fragramce.
 - a surfactant,
 - an antimicrobial agent,
 - a corrosion inhibitor, and/or
 - a miticide.

09/509308 430 Rec'd PCT/PTO 23 MAR 2000

Abstract of the Disclosure

Der-f and/or Der-p dust mite allergens are deactivated by an amount of one or more of the following deactivants: i) cedarwood oil, ii) hexadecyltrimethylammonium chloride, iii) aluminium chlorohydrate, iv) 1-propoxy-propanol-2, v) polyquaternium-10 vi) silica gel, vii) propylene glycol alginate, viii) ammonium sulphate, ix) hinokitiol, x) L-ascorbic acid, xi) immobilised tannic acid, xii) chlorohexidine, xiii) maleic anhydride, xiv) hinoki oil, xv) a composite of AgCl and TiO₂, xvi) diazolidinyl urea, xvii) 6-isopropyl-m-cresol, xviii) a compound of formula I

$$O$$
 octyl O octyl O octyl O

10 xix) the compound of formula II

$$O \longrightarrow CH_3$$
 $O \longrightarrow CH_2$
 $O \longrightarrow CH_3$

xx) a polymeric dialdehyde containing two or more of a recurring unit of the formula III

where n = 2 to 200, xxi) urea, xxii) cyclodextrin, xxiii) hydrogenated hop oil, xxiv) polyvinylpyrrolidone, xxv) N-methylpyrrolidone, xxvi) the sodium salt of
anthraquinone, xxvii) potassium thioglycolate, and xxviii) glutaraldehyde.
Deactivants (i) to (xx) are effective against allergens derived from both species.
Deactivants (xxi) to (xxvi) are effective against only Der-f allergens. Deactivants (xxvii) and (xxviii) are effective against only Der-p allergens. Aerosol compositions comprise said deactivants, a propellant and optional solvents.

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DOCKET NO: 08291-435001

COMBINED DECLARATION AND POWER OF ATTORNEY FOR UTILITY PATENT APPLICATION

As a below named inventor, I hereby declare that:

My residence, post office address and citizenship are as stated below next to my name,

I believe I am the original, first and sole inventor (if only one name is listed below) or an original, first and joint inventor (if plural names are listed below) of the subject matter which is claimed and for which a patent is sought on the invention entitled Deactivants for Dust Mite Allergens the specification of which is attached hereto. as Application Serial mas filed on _____ No. _____ and was amended was described and claimed in PCT International Application No. PCT/GB98/02863 filed on 22 September 1998 and was amended under PCT Article 19 on _ I hereby state that I have reviewed and understand the contents of the aboveidentified specification, including the claims, as amended by any amendment referred to above. I acknowledge the duty to disclose all information I know to be material to patentability in accordance with 37 C.F.R. § 1.56. I hereby claim foreign priority benefits under 35 U.S.C. § 119 of any foreign application(s) for patent or inventor's certificate or of any PCT International application(s) designating at least one country other than the United States of America listed below and have also identified below any foreign application for patent or inventor's certificate or any PCT international application(s) designating at least one country other than the United States of America filed by me on the same subject matter having a filing date before that of the application(s) of which priority is claimed: PRIORITY CLAIMED APPLICATION NO. FILING DATE **COUNTRY** 25 September 1997 X Yes D No 9720275.8 25 September 1997 X Yes No 9720298.0 GB I hereby claim the benefit under 35 U.S.C. § 120 of any United States

I hereby claim the benefit under 35 U.S.C. § 120 of any United States application(s) or under 35 U.S.C. § 365(c) of any PCT International application designating the United States of America listed below and, insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States or PCT International application in the manner provided by the first paragraph of 35 U.S.C. § 112, I acknowledge the duty to disclose all information I know to be material to patentability as defined in 37 C.F.R. § 1.56(a) which became available between the filing date of the prior application and the national or PCT International filing date of this application:

COMBINED DECLARATION AND POWER OF ATTORNEY (CONTINUED)

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APPLICATION NO.	FILING DATE	U.S. PATEN ☐ Pending	IT STATUS ☐ Issued ☐ Abandoned
I hereby appoint the and to transact all business Frederick H. Rabin, Reg. N. Pegram, Reg. No. 25,198; V. Stephan J. Filipek, Reg. No.	in the Patent and Tra o_24,488; Andrew N William J. Hone, Reg	demark Office . Parfomak, Ro . No. 26,739; I	eg. No. <u>32,431;</u> John B. Richard P. Ferrara, 30,632;
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that all statements made on these statements were made	information and belic with the knowledge fine or imprisonment,	ef are believed that willful fal or both, unde	se statements and the like at 18 U.S.C. § 1001 and that
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